Serial No.: 10/511,993 Filed: March 28, 2005

Page : 2 of 10

Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

1. (Currently amended) A method of screening for agonistic antibodies that comprises

the following steps (a) to (c):

(a) providing a cell that expresses both a multimer-forming receptor and a test antibody,

wherein the cell grow depending on the corresponding ligand of the receptor;

(b) determining the test antibody to comprise agonistic activity when autocrine cell

growth is autonomous; and

(c) selecting those antibodies as that comprise agonistic activity.

2. (Original) The method of claim 1, that further comprises the step of introducing a

gene that encodes the heavy chain of the test antibody, into the cell of step (a) having been

introduced with a gene that encodes the light chain of the test antibody, and a gene that encodes

the receptor.

3. (Previously presented) The method of claim 1 wherein the receptor is a chimeric

receptor with a protein that comprises a function of transducing a cell growth signal.

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 3 of 10

4. (Previously presented) The method of claim 1 wherein the receptor is a dimer-

forming receptor.

5. (Previously presented) The method of claim 4 wherein the dimer-forming receptor is

a homo-dimer.

6. (Previously presented) The method of claim 4 wherein the dimer-forming receptor is

a hetero-dimer.

7. (Previously presented) The method of claim 1 wherein the protein that comprises the

function of transducing a cell growth signal is a G-CSF receptor.

8. (Previously presented) The method of claim 1 that comprises the introduction of an

antibody library to the cell.

9. (Previously presented) The method of claim 8 wherein the antibody library is a

retroviral antibody library.

10. (Previously presented) The method of claim 1 wherein the test antibody is a multi-

specific antibody.

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 4 of 10

11. (Original) The method of claim 10 that comprises linking the test antibody's heavy

and light chain variable regions with a linker.

12. (Original) The method of claim 11 that comprises producing the antibody with

variable regions linked by a linker, using a method that comprises the steps (a) to (c):

(a) producing a single chain Fv against the first receptor chain;

(b) producing a single chain antibody against the first receptor chain by linking the single

chain Fv with a CH1-hinge-CH2-CH3; and

(c) producing a multi-specific antibody that comprises the single chain antibody produced

in step (b).

13. (Original) The method of claim 11 that comprises producing the antibody with its

variable regions linked by a linker, using a method that comprises the steps (a) to (c):

(a) producing a single chain Fab against the first receptor chain;

(b) producing a single chain antibody against the first receptor chain by linking the single

chain Fab with an Fc; and

(c) producing a multi-specific antibody that comprises the single chain antibody produced

in step (b).

14. (Withdrawn) A method of screening for an agonist multi-specific antibody that

comprises the steps (a) to (c):

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 5 of 10

(a) contacting between a multi-specific antibody and a receptor comprising a first

receptor chain and a second receptor chain, where the multi-specific antibody comprises a

variable region that can bind with the first receptor chain and a variable region that can bind with

the second receptor chain;

(b) determining whether the test multi-specific antibody comprises agonistic activity; and

(c) selecting antibodies that comprise agonistic activity.

15. (Withdrawn) The method of claim 14 that comprises expressing the receptor and the

test multi-specific antibody in the same cell.

16. (Withdrawn) The method of claim 15 wherein the cell is a cell that grows depending

on the corresponding ligand of the receptor.

17. (Withdrawn) The method of claim 15 wherein the receptor comprises the function of

transducing a cell growth signal.

18. (Withdrawn) The method of claim 17 wherein the receptor is a chimeric receptor

with a protein that comprises the function of transducing a cell growth signal.

19. (Withdrawn) The method of claim 18 wherein the protein that comprises the

function of transducing a cell growth signal is a G-CSF receptor.

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 6 of 10

20. (Withdrawn) The method of claim 15 wherein the test multi-specific antibody is

determined to comprise agonistic activity when autocrine cell growth is autonomous.

21. (Withdrawn) The method of claim 15 that further comprises the step of introducing

an antibody library against the first receptor chain and the second receptor chain into the cell,

respectively.

22. (Withdrawn) The method of claim 21 wherein the antibody library is a retroviral

antibody library.

23. (Withdrawn) The method of claim 14 that comprises linking the light chain variable

regions and heavy chain variable regions of the multi-specific antibody with a linker.

24. (Withdrawn) The method of claim 23 that comprises producing a multi-specific

antibody with variable regions linked by a linker, using a method that comprises steps (a) to (c):

(a) producing a single chain Fv against the first receptor chain;

(b) producing a single chain antibody against the first receptor chain by linking the single

chain Fv with a CH1-hinge-CH2-CH3; and

(c) producing a multi-specific antibody that comprises the single chain antibody produced

in step (b).

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 7 of 10

25. (Withdrawn) The method of claim 23 that comprises producing the multi-specific

antibody with variable regions linked by a linker, using a method that comprises steps (a) to (c):

(a) producing a single chain Fab against the first receptor chain;

(b) producing a single chain antibody against the first receptor chain by linking the single

chain Fab with an Fc; and

(c) producing a multi-specific antibody that comprises the single chain antibody produced

in step (b).

26. (Withdrawn) The method claim 14 that comprises the introduction of "Knobs-into-

holes" by amino acid substitution at the CH3 region of the multi-specific antibody.

27. (Withdrawn) The method claim 14 wherein the multimer of the receptor is a

heterodimer.

28. (Withdrawn) The method claim 14 wherein the multi-specific antibody is a

bispecific antibody.

29. (Withdrawn) A method for producing an agonistic antibody comprising steps (a) to

(c):

(a) screening for an agonistic antibody by a method claim 1;

Serial No.: 10/511,993 Filed: March 28, 2005

Page : 8 of 10

(b) introducing a gene that encodes the agonistic antibody selected by the screening of

step (a) into a host cell;

(c) recovering the agonistic antibody from the host cell of step (b) or its cell culture

supernatant.

30. (Withdrawn) A cell that expresses an antibody, and a receptor that multimerizes by

binding with the antibody, where the cell grow depending on the corresponding ligand of the

receptor.

31. (Withdrawn) The cell of claim 30 where the receptor is a chimeric receptor with a

protein that comprises the function of transducing a cell growth signal.

32. (Withdrawn) The cell of claim 30 wherein the antibody is a multi-specific antibody.

33. (Withdrawn) The cell of claim 30 wherein the receptor that is multimerized by

binding with the antibody comprises the function of transducing a cell growth signal.

34. (Withdrawn) A multi-specific agonistic antibody that comprises the linking of the

light chain variable region and heavy chain variable region by linkers, and the introduction of

"Knobs-into-holes" by amino acid substitution at the CH3 region of the antibody.